WHAT IS CLAIMED:

1. A compound of Formula I:

5

wherein

A is selected from:

(a) hydrogen;

(b) $-(C=O)-O-R_1$, where R_1 is selected from:

10

- 1. hydrogen,
- 2. C_1-C_6 alkyl,
- 3. C₃-C₁₂ cycloalkyl,
- 4. substituted C₃-C₁₂ cycloalkyl,
- 5. aryl,
- 6. substituted aryl,
- 7. heteroaryl,
- 8. substituted heteroaryl,
- 9. heterocycloalkyl,

10. substituted heterocycloalkyl, or

20

15

- 11.–C₁–C₆ alkyl containing 0, 1, 2, or 3 heteroatoms selected from O, S, or N, optionally substituted with one or more substituent selected from halogen, aryl, substituted aryl, heteroaryl, substituted heteroaryl, heterocycloalkyl, or substituted heterocycloalkyl;
- (c) $-(C=O)-R_2$, where R_2 is selected from:

- 1. -R₁, where R₁ is as previously defined,
- 2. alkylamino,

- 3. dialkyl amino,
- 4. arylamino, or
- 5. diarylamino;
- (d) $-C(=O)-NH-R_2$, where R_2 is as previously defined;
- (e) -C(=S)-NH-R₂, where R₂ is as previously defined;
- (f) $-S(O)_2-R_2$, where R_2 is as previously defined;

B is hydrogen or C₁–C₆ alkyl;

G is

5

15

- (a) -OH;
- 10 (b) $-O-(C_1-C_{12} \text{ alkyl});$
 - (c) -NH-R₂, where R₂ is as previously defined;
 - (d) $-NHS(O)_2-R_1$, where R_1 as previously defined;
 - (e) $-(C=O)-R_2$, where R_2 as previously defined;
 - (f) -(C=O)-O-R₁, where R₁ as previously defined; or
 - (g) $-(C=O)-NH-R_2$, where R_2 as previously defined;

M is absent or selected from:

- (a) -0-;
- (b) -S-;
- (c) -NH-; or
- 20 (d) $-NR_1$ -, wherein R_1 is previously defined;

Q is selected from:

- (a) aryl;
- (b) substituted aryl;
- (c) heteroaryl;
- 25 (d) substituted heteroaryl;
 - (e) heterocycloalkyl; or
 - (f) substituted heterocycloalkyl;

j = 0, 1, 2, 3, or 4;

n = 0, 1, or 2; and

30 s = 0, 1, or 2.



2. A compound of formula I, wherein M is absent and Q is

10

15

30

wherein X and Y are each independently selected from:

- a) -C₁-C₆ alkyl containing 0, 1, 2, or 3 heteroatoms selected from O, S, or N, optionally substituted with one or more substituent selected from halogen, aryl, substituted aryl, heteroaryl, substituted heteroaryl, heterocycloalkyl, or substituted heterocycloalkyl;
- b) -C₂-C₆ alkenyl containing 0, 1, 2, or 3 heteroatoms selected from O, S, or N, optionally substituted with one or more substituent selected from halogen, aryl, substituted aryl, heteroaryl, substituted heteroaryl, heterocycloalkyl, or substituted heterocycloalkyl;
- c) -C₂-C₆ alkynyl containing 0, 1, 2, or 3 heteroatoms selected from O, S, or N, optionally substituted with one or more substituent selected from halogen, aryl, substituted aryl, heteroaryl, substituted heteroaryl, heterocycloalkyl, or substituted heterocycloalkyl;
- d) aryl;
- e) substituted aryl;
- f) heteroaryl;
- g) substituted heteroaryl;
- h) heterocycloalkyl; or
- substituted heterocycloalkyl;
- or in the alternative, X and Y are taken together with the carbons to which they are attached to for a cyclic moiety selected from: aryl, substituted aryl, heteroaryl, substituted heteroaryl, heterocycloalkyl, or substituted heterocycloalkyl;



- 25 3. A compound of formula I, wherein M is absent and Q is wherein Y is selected from:
 - a) -C₁-C₆ alkyl containing 0, 1, 2, or 3 heteroatoms selected from O, S, or N, optionally substituted with one or more substituent selected from halogen, aryl, substituted aryl, heteroaryl, substituted heteroaryl, heterocycloalkyl, or substituted heterocycloalkyl;

- b) -C₂-C₆ alkenyl containing 0, 1, 2, or 3 heteroatoms selected from O, S, or N, optionally substituted with one or more substituent selected from halogen, aryl, substituted aryl, heteroaryl, substituted heteroaryl, heterocycloalkyl, or substituted heterocycloalkyl;
- c) -C₂-C₆ alkynyl containing 0, 1, 2, or 3 heteroatoms selected from O, S, or N, optionally substituted with one or more substituent selected from halogen, aryl, substituted aryl, heteroaryl, substituted heteroaryl, heterocycloalkyl, or substituted heterocycloalkyl;
- d) aryl;
- e) substituted aryl;
- f) heteroaryl;
- g) substituted heteroaryl;
- h) heterocycloalkyl; or
- i) substituted heterocycloalkyl;

20

25

10

5

- 4. A compound of formula I, wherein M is absent and Q is wherein X, Y, and Z are each independently selected from:
 - a) -C₁-C₆ alkyl containing 0, 1, 2, or 3 heteroatoms selected from O, S, or N, optionally substituted with one or more substituent selected from halogen, aryl, substituted aryl, heteroaryl, substituted heteroaryl, heterocycloalkyl, or substituted heterocycloalkyl;
 - b) -C₂-C₆ alkenyl containing 0, 1, 2, or 3 heteroatoms selected from O, S, or N, optionally substituted with one or more substituent selected from halogen, aryl, substituted aryl, heteroaryl, substituted heteroaryl, heterocycloalkyl, or substituted heterocycloalkyl;
 - c) -C₂-C₆ alkynyl containing 0, 1, 2, or 3 heteroatoms selected from O, S, or N, optionally substituted with one or more substituent selected from halogen, aryl, substituted aryl, heteroaryl, substituted heteroaryl, heterocycloalkyl, or substituted heterocycloalkyl;

- d) aryl;
- e) substituted aryl;

15

20

25

- f) heteroaryl;
- g) substituted heteroaryl;
- h) heterocycloalkyl; or
- i) substituted heterocycloalkyl;

or in the alternative, X and Y or Y and Z are taken together with the carbons to which they are attached to for a cyclic moiety selected from: aryl, substituted aryl, heteroaryl, substituted heteroaryl, heterocycloalkyl, or substituted heterocycloalkyl;

10 5. A compound of formula I, wherein M is absent and Q is

wherein W, X, Y, and Z are each independently selected from:

- a) -C₁-C₆ alkyl containing 0, 1, 2, or 3 heteroatoms selected from O, S, or N, optionally substituted with one or more substituent selected from halogen, aryl, substituted aryl, heteroaryl, substituted heteroaryl, heterocycloalkyl, or substituted heterocycloalkyl;
- b) -C₂-C₆ alkenyl containing 0, 1, 2, or 3 heteroatoms selected from O, S, or N, optionally substituted with one or more substituent selected from halogen, aryl, substituted aryl, heteroaryl, substituted heteroaryl, heterocycloalkyl, or substituted heterocycloalkyl;
- c) -C₂-C₆ alkynyl containing 0, 1, 2, or 3 heteroatoms selected from O, S, or N, optionally substituted with one or more substituent selected from halogen, aryl, substituted aryl, heteroaryl, substituted heteroaryl, heterocycloalkyl, or substituted heterocycloalkyl;
- d) aryl;
- e) substituted aryl;
- f) heteroaryl;
- g) substituted heteroaryl;
- h) heterocycloalkyl; or
- i) substituted heterocycloalkyl:
- or in the alternative, W and X, X and Y, or Y and Z are taken together with the carbons to which they are attached to for a cyclic moiety selected from:

10

15

20

25

aryl, substituted aryl, heteroaryl, substituted heteroaryl, heterocycloalkyl, or substituted heterocycloalkyl;

$$x \stackrel{N}{\swarrow}_{N}^{Y} z$$

- 6. A compound of formula I, wherein M is absent and Q is wherein X, Y, and Z are each independently selected from:
 - a) -C₁-C₆ alkyl containing 0, 1, 2, or 3 heteroatoms selected from O, S, or N, optionally substituted with one or more substituent selected from halogen, aryl, substituted aryl, heteroaryl, substituted heteroaryl, heterocycloalkyl, or substituted heterocycloalkyl;
 - b) -C₂-C₆ alkenyl containing 0, 1, 2, or 3 heteroatoms selected from O, S, or N, optionally substituted with one or more substituent selected from halogen, aryl, substituted aryl, heteroaryl, substituted heteroaryl, heterocycloalkyl, or substituted heterocycloalkyl;
 - c) $-C_2-C_6$ alkynyl containing 0, 1, 2, or 3 heteroatoms selected from O, S, or N, optionally substituted with one or more substituent selected from halogen, aryl, substituted aryl, heteroaryl, substituted heteroaryl, heterocycloalkyl, or substituted heterocycloalkyl;
 - d) aryl;
 - e) substituted aryl;
 - f) heteroaryl;
 - g) substituted heteroaryl;
 - h) heterocycloalkyl; or
 - i) substituted heterocycloalkyl;
 - or in the alternative, Y and Z are taken together with the carbons to which they are attached to for a cyclic moiety selected from: aryl, substituted aryl, heteroaryl, substituted heteroaryl, heterocycloalkyl, or substituted heterocycloalkyl;

7. A compound of formula I, wherein M is -O- and Q is

wherein

5

10

15

20

25

L is M, where M is as previously defined;

X, Y, and Z are each independently selected from:

- a) -C₁-C₆ alkyl containing 0, 1, 2, or 3 heteroatoms selected from O, S, or N, optionally substituted with one or more substituent selected from halogen, aryl, substituted aryl, heteroaryl, substituted heteroaryl, heterocycloalkyl, or substituted heterocycloalkyl;
- b) -C₂-C₆ alkenyl containing 0, 1, 2, or 3 heteroatoms selected from O, S, or N, optionally substituted with one or more substituent selected from halogen, aryl, substituted aryl, heteroaryl, substituted heteroaryl, heterocycloalkyl, or substituted heterocycloalkyl;
- c) -C₂-C₆ alkynyl containing 0, 1, 2, or 3 heteroatoms selected from O, S, or N, optionally substituted with one or more substituent selected from halogen, aryl, substituted aryl, heteroaryl, substituted heteroaryl, heterocycloalkyl, or substituted heterocycloalkyl;
- d) aryl;
- e) substituted aryl;
- f) heteroaryl;
- g) substituted heteroaryl;
- h) heterocycloalkyl; or
- i) substituted heterocycloalkyl;
- or in the alternative, X and Y are taken together with the carbons to which they are attached to for a cyclic moiety selected from: aryl, substituted aryl, heteroaryl, substituted heteroaryl, heterocycloalkyl, or substituted heterocycloalkyl;

8. A compound of formula I, wherein M is -O- and Q is wherein

L is M, where M is as previously defined;

30 X, Y, and Z are each independently selected from:

10

15

20

25

30

- a) -C₁-C₆ alkyl containing 0, 1, 2, or 3 heteroatoms selected from O, S, or N, optionally substituted with one or more substituent selected from halogen, aryl, substituted aryl, heteroaryl, substituted heteroaryl, heterocycloalkyl, or substituted heterocycloalkyl;
- b) -C₂-C₆ alkenyl containing 0, 1, 2, or 3 heteroatoms selected from O, S, or N, optionally substituted with one or more substituent selected from halogen, aryl, substituted aryl, heteroaryl, substituted heteroaryl, heterocycloalkyl, or substituted heterocycloalkyl;
- c) -C₂-C₆ alkynyl containing 0, 1, 2, or 3 heteroatoms selected from O, S, or N, optionally substituted with one or more substituent selected from halogen, aryl, substituted aryl, heteroaryl, substituted heteroaryl, heterocycloalkyl, or substituted heterocycloalkyl;
- d) aryl;
- e) substituted aryl;
- f) heteroaryl;
- g) substituted heteroaryl;
- h) heterocycloalkyl; or
- i) substituted heterocycloalkyl;

or in the alternative, X and Y are taken together with the carbons to which they are attached to for a cyclic moiety selected from: aryl, substituted aryl, heteroaryl, substituted heteroaryl, heterocycloalkyl, or substituted heterocycloalkyl; or

is x L Z

9. A compound of formula I, wherein M is -O- and Q is

wherein

L is M, where M is as previously defined;

- X, Y, and Z are each independently selected from:
- a) -C₁-C₆ alkyl containing 0, 1, 2, or 3 heteroatoms selected from O, S, or N, optionally substituted with one or more substituent selected from halogen, aryl, substituted aryl, heteroaryl, substituted heteroaryl, heterocycloalkyl, or substituted heterocycloalkyl;

- b) -C₂-C₆ alkenyl containing 0, 1, 2, or 3 heteroatoms selected from O, S, or N, optionally substituted with one or more substituent selected from halogen, aryl, substituted aryl, heteroaryl, substituted heteroaryl, heterocycloalkyl, or substituted heterocycloalkyl;
- c) -C₂-C₆ alkynyl containing 0, 1, 2, or 3 heteroatoms selected from O, S, or N, optionally substituted with one or more substituent selected from halogen, aryl, substituted aryl, heteroaryl, substituted heteroaryl, heterocycloalkyl, or substituted heterocycloalkyl;
- d) aryl;
- 10 e) substituted aryl;
 - f) heteroaryl;
 - g) substituted heteroaryl;
 - h) heterocycloalkyl; or
 - i) substituted heterocycloalkyl;
- or in the alternative, X and Y are taken together with the carbons to which they are attached to for a cyclic moiety selected from: aryl, substituted aryl, heteroaryl, substituted heteroaryl, heterocycloalkyl, or substituted heterocycloalkyl.
- 10. A compound according to claim 1 represented by formula I selected from: Compound of formula I, wherein A = Boc, B = hydrogen, G = OEt, M = -O-, Q = hydrogen, and j = n = s = 1;
- Compound of formula I, wherein A = Boc, B = hydrogen, G = OEt, M = -O-, Q = $-S(O)_2CH_3$, and j = n = s = 1;
 - Compound of formula I, wherein A = Boc, B = hydrogen, G = OEt, M = -O-, Q = 2-thiophenyl-quinolin-4-yl, and j = n = s = 1;
- Compound of formula I, wherein A = Boc, B = hydrogen, G = OH, M = -O-, Q = 2-thiophenyl-quinolin-4-yl, and j = n = s = 1;

- Compound of formula I, wherein A = Boc, B = hydrogen, G = OEt, M is absent, Q = 4,5-diphenyltriazol-2-yl, and j = n = s = 1;
- Compound of formula I, wherein A = Boc, B = hydrogen, G = OH, M is absent, Q = 4,5-di-thiophenyltriazol-2-yI, and j = n = s = 1;
 - Compound of formula I, wherein A = Boc, B = hydrogen, G = OH, M is absent, Q = 4-(thiophen-3-yl)-5-(p-methoxyphenyl)triazol-2-yl, and <math>j = n = s = 1;
- 10 Compound of formula I, wherein A = Boc, B = hydrogen, G = OH, M is absent, Q = 4-(n-butyl)-5-phenyltriazol-2-yl, and j = n = s = 1;
 - Compound of formula I, wherein A = Boc, B = hydrogen, G = OH, M is absent, Q = 5-(3-methoxyphenyl)tetrazol-2-yl, and j = n = s = 1;
 - Compound of formula I, wherein A = Boc, B = hydrogen, G = OH, M is absent, Q = 5-(4-pyridyI)tetrazol-2-yI, and j = n = s = 1;
- Compound of formula I, wherein A = Boc, B = hydrogen, G = OH, M is absent, Q = 5-(3,4-dichlorophenyl)tetrazol-2-yl, and j = n = s = 1;
 - Compound of formula I, wherein A = Boc, B = hydrogen, G = OH, M is absent, Q = 5-(3-bromo-4-methoxy-phenyl)tetrazol-2-yl, and j = n = s = 1;
- Compound of formula I, wherein A = Boc, B = hydrogen, G = OH, M is absent, Q = 4-(4-fluoro-phenyl)-6-phenyl-1H-pyridazin-3-on-2-yl, and j = n = s = 1;
 - Compound of formula I, wherein A = Boc, B = hydrogen, G = OH, M is absent, Q = 6-phenyl-5-piperidin-1-yl-1H-pyridazin-3-on-2-yl, and j = n = s = 1;
 - Compound of formula I, wherein A = Boc, B = hydrogen, G = OH, M = -O-, Q = 7-Methoxy-2-phenyl-quinolin-4-yl, and j = n = s = 1;

WO 2005/010029 PCT/US2004/015802 68

- Compound of formula I, wherein A = Boc, B = hydrogen, G = OH, M = -O-, Q =7-Methoxy-2-thiazolyl-quinolin-4-yl, and j = n = s = 1;
- Compound of formula I, wherein A = Boc, B = hydrogen, G = OH, M = -O-, Q =7-Methoxy-2-thiophenyl-quinolin-4-yl, and i = n = s = 1;

5

- Compound of formula I, wherein A = Boc, B = hydrogen, G = OH, M = -O-, Q =7-Methoxy-3-(thiophen-2-yl)-1H-quinoxalin-2-yl, and j = n = s = 1;
- 10 Compound of formula I, wherein A = Boc, B = hydrogen, G = OH, M = -O-, Q =6-Methoxy-3-(thiophen-2-vI)-1H-quinoxalin-2-vI, and i = n = s = 1:
 - Compound of formula I, wherein A = Boc, B = hydrogen, G = OH, M = -O-, Q =7-Methoxy-3-[2-(thiophen-2-yl)vinyl]-1H-quinoxalin-2-yl, and j = n = s = 1;
 - Compound of formula I, wherein A = Boc, B = hydrogen, G = OH, M = -O-, Q = 6-Methoxy-3-[2-(thiophen-2-yl)vinyl]-1H-quinoxalin-2-yl, and j = n = s = 1;
- Compound of formula I, wherein A = Boc, B = hydrogen, G = OH, M = -O-, Q =20 7-Methoxy-3-[2-(pyridin-2-yl)vinyl]-1H-quinoxalin-2-yl, and j = n = s = 1; or
 - Compound of formula I, wherein A = Boc, B = hydrogen, G = OH, M = -O-, Q = 7-methoxy-3-[2-(pyridin -2-yl)vinyl]-1H-quinoxalin-2-yl, and j = n = s = 1.
- 25 11. A pharmaceutical composition comprising an anti-hepatitis C virally effective amount of a compound according to claim 1, or a pharmaceutically acceptable salt, ester, or prodrug thereof, in combination with a pharmaceutically acceptable carrier or excipient.
- 30 12. A method of treating a hepatitis C viral infection in a mammal, comprising administering to the mammal an anti-hepatitis C virally effective amount of a pharmaceutical composition according to claim 11.

15

20

- 13. A method of inhibiting the replication of hepatitis C virus, the method comprising supplying a hepatitis C viral NS3 protease inhibitory amount of the pharmaceutical composition of claim 11.
- 5 14. The method of claim 13 further comprising administering concurrently an additional anti-hepatitis C virus agent.
 - 15. The method of claim 14, wherein said additional anti-hepatitis C virus agent is selected from the group consisting of: α -interferon, β -interferon, ribavarin, and adamantine.
 - 16. The method of claim 14, wherein said additional anti-hepatitis C virus agent is an inhibitor of another target in the hepatitis C virus life cycle, which is selected from the group consisting of: helicase, polymerase, metalloprotease, and IRES.
 - 17. A process of making compounds of formula I:

wherein A, B, G, M, Q, j, n, and s are as defined in claim 1, comprising the steps of:

10

wherein A, B, and j are as defined in claim 1 and T is selected from OH, OMe, or OEt;

(b) reacting a compound of formula B with a compound of formula (D):

, wherein G is as defined in claim 1, under standard amide formation conditions to form a compound of formula (**E**):

, wherein A, B, G, M, Q, and j are as defined in

claim 1; and

(c) reacting compound of formula E with a Ruthenium-based catalyst.